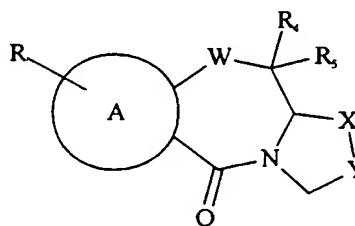
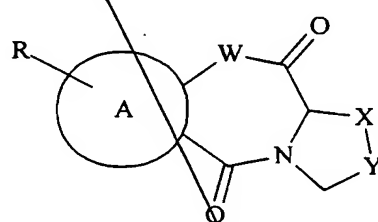


We claim:

1. A method of inhibiting a HIV integrase, the method comprising:
 exposing the integrase to an integrase inhibiting amount of one or more
 anti-integrase compounds selected from the group consisting of the following
 5 compounds, or pharmaceutically acceptable salts thereof:



wherein

A is thiazole, benzene, naphthalene, pyridine, pyrimidine, pyrazine, or quinoline;

R is one or more of H, halogen, lower alkyl, lower alkoxy, NO₂, lower ester or carboxylic acid;

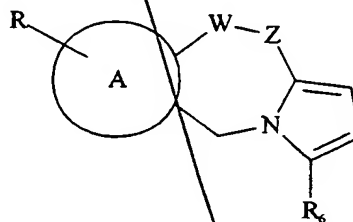
X-Y is CH₂-S, S-CH₂, CH₂-O, CH₂-S(O), S(O)-CH₂, CH₂-CH₂, CH₂-CH₂-CH₂, or CH₂-CH₂-CH₂-CH₂;

R₄ is H or hydroxy;

R₅ is H, phenyl, or alkylamine; and

W is S or O.

or wherein the compound is



wherein

A is thiazole, benzene, naphthalene, pyridine, pyrimidine, pyrazine, or quinoline; and

R is one or more of H, halogen, lower alkyl, lower ester or carboxylic acid;

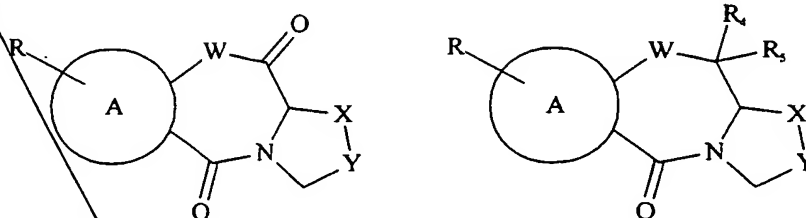
R₆ is H, substituted or unsubstituted alkyl or amine;

W is S or O; and

Z is S, O, CH₂, CH₂CH₂, or C=O.

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2. The method of claim 1, wherein the compound is selected from the group consisting of:



wherein X-Y is CH₂-S, S-CH₂, CH₂-O, or CH₂-CH₂, and W is S.

3. The method of claim 2, wherein:

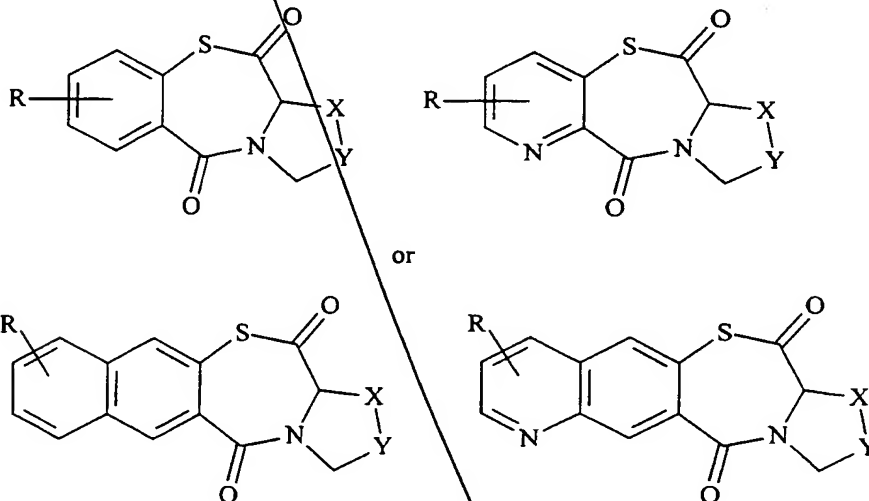
A is benzene, naphthalene, pyridine, pyrimidine, pyrazine, or quinoline.

4. The method of claim 3, wherein A is benzene or naphthalene.

5. The method of claim 4, wherein R is H, halogen, lower alkoxy, or

NO₂.

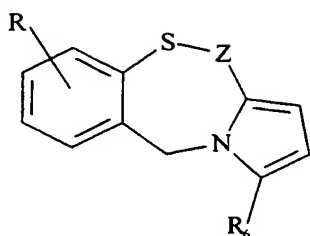
6. The method of claim 1, wherein the compound is:



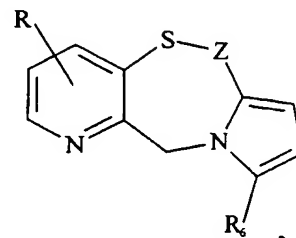
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- 40 -

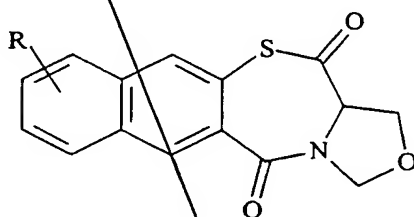
7. The method of claim 1, wherein the compound is:



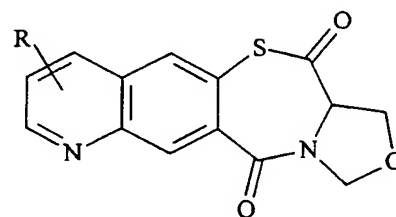
or



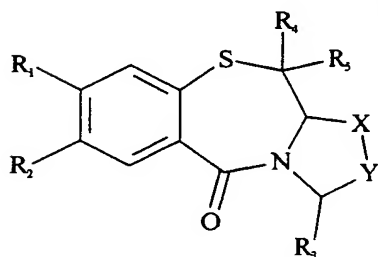
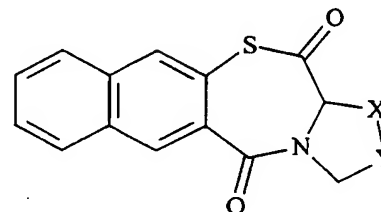
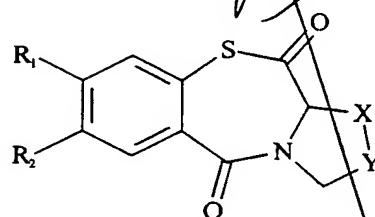
8. The method of claim 6, wherein the compound is



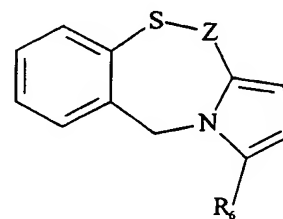
or



- 5 9. The method of claim 1, wherein the compound is one of the following:



or



wherein X-Y is S-CH₂, CH₂-S, CH₂-O, CH₂-CH₂, S(O)-CH₂, or CH₂-S(O);

R₁ and R₂ are independently selected from the group consisting of H, NO₂,

10 halogen, lower alkyl or lower alkoxy;

R₃ is H or phenyl;

R₄ is H or hydroxy;

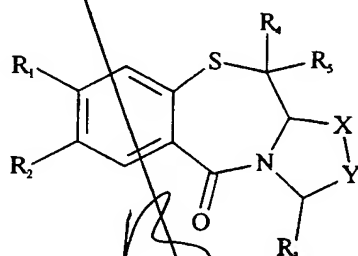
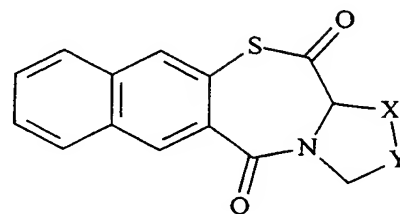
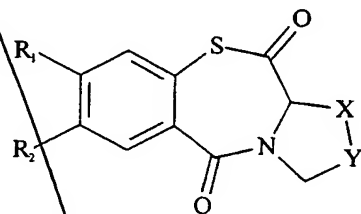
R₅ is H, phenyl or alkylamine; and

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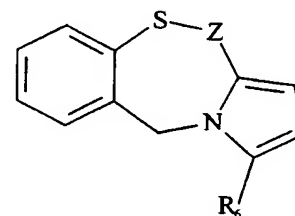
R_6 is H, phenyl or alkylamine.

10. The method of claim 9, wherein the alkylamine is $-N(CH_2CH_2)_2NCH_3$, $-CH_2NCH_2CH_3$, or $-CH_2N(CH_2CH_2)_2NCH_3$.

11. The method of claim 7, wherein the compound is



or



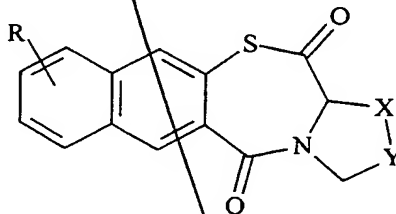
X-Y is $S-CH_2$, CH_2-S , or $CH_2-S(O)$;

and R_1 and R_2 are independently selected from the group consisting of H, NO_2 , halogen, lower alkyl and lower alkoxy;

R_3 is H; and

R_4 , R_5 , and R_6 are H.

12. The method of claim 1, wherein the compound is



and X-Y is $S-CH_2$ or CH_2-S .

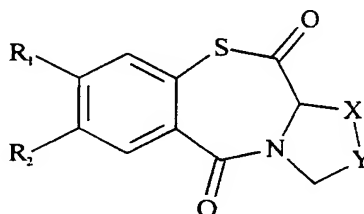
13. The method of claim 12, wherein R is H.

14. The method of claim 13, wherein X-Y is $S-CH_2$.

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15. The method of claim 9, wherein
R₁ is H, NO₂, or lower alkoxy,
R₂ is H, Cl, Br, lower alkyl, or lower alkoxy;
R₃ and R₄ are H;
R₅ is N(CH₂CH₂)₂NCH₃; and
X-Y is CH₂-S, S-CH₂, or CH₂-CH₂.

16. The method of claim 15 wherein the compound is



- wherein R₁ is H, NO₂, or methoxy;
R₂ is H, halogen or methoxy; and
X-Y is CH₂-S or S-CH₂.

17. The method of claim 1, wherein the compound is administered in a therapeutically effective amount to a subject.

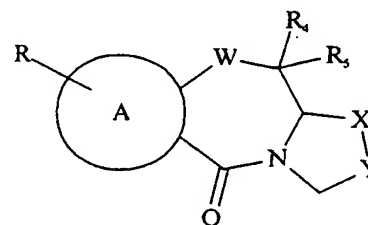
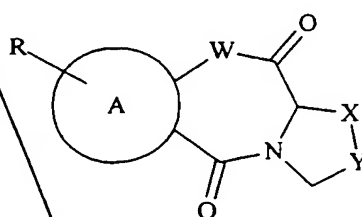
18. The method of claim 17, wherein the method is a method of treating or preventing HIV infection in the subject.

19. The method of claim 15, wherein the compound is administered in a therapeutically effective amount to a subject to treat or prevent an HIV infection.

20. The method of claim 16, wherein the compound is administered in a therapeutically effective amount to a subject to treat or prevent an HIV infection.

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21. A method of treating or preventing HIV infection in a subject, comprising administering to the subject a therapeutically effective amount of a compound selected from the group consisting of:



wherein

5 A is thiazole, benzene, naphthalene, pyridine, pyrimidine, pyrazine, or quinoline;

R is one or more of H, halogen, lower alkyl, lower alkoxy, NO₂, lower ester or carboxylic acid;

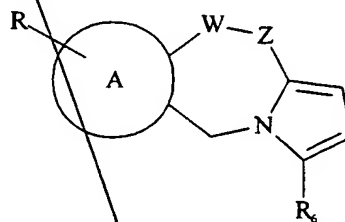
10 X-Y is CH₂-S, S-CH₂, CH₂-O, CH₂-S(O), S(O)-CH₂, CH₂-CH₂, CH₂-CH₂-CH₂, or CH₂-CH₂-CH₂-CH₂;

R₄ is H or hydroxy;

R₅ is H, phenyl, or alkylamine; and

W is S or O.

15 or wherein the compound is



wherein

A is thiazole, benzene, naphthalene, pyridine, pyrimidine, pyrazine, or quinoline; and

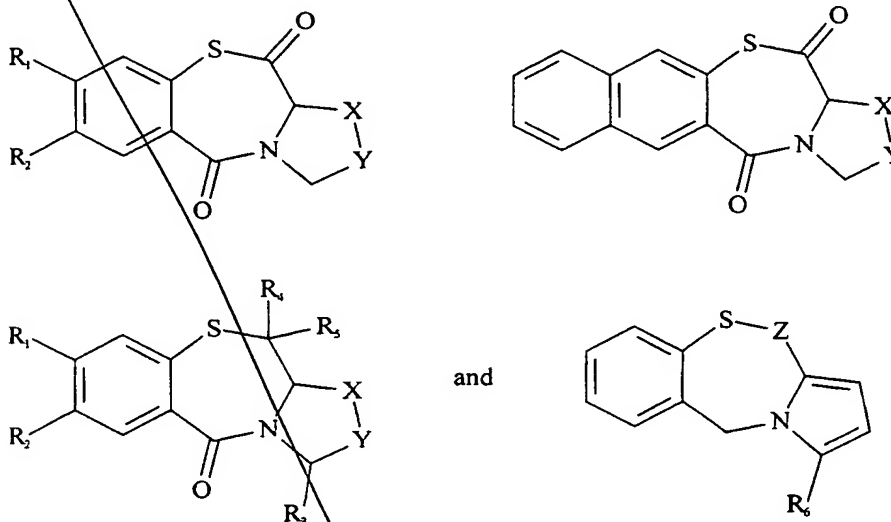
20 R is selected from the group of H, halogen, lower alkyl, lower ester or carboxylic acid;

R₆ is H, substituted or unsubstituted alkyl or amine;

W is S or O; and

Z is S, O, CH₂, CH₂CH₂, or C=O.

22. The method of claim 21, wherein the compound is selected from the group consisting of:



wherein X-Y is S-CH₂, CH₂-S, CH₂CH₂ or S(O)CH₂;

R₁ is H, NO₂, or lower alkoxy;

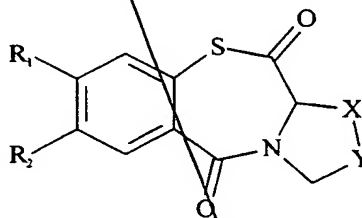
R₂ is H, Cl, Br, lower alkyl, or lower alkoxy;

R₃ and R₄ are H;

R₅ is N(CH₂CH₂)₂NCH₃; and

R₆ is H.

23. The method of claim 21, wherein the compound is



and

R₁ and R₂ are H, and X-Y is S-CH₂; or

R₁ is H, R₂ is Cl or Br or methyl, and X-Y is S-CH₂; or

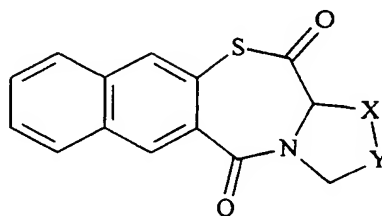
R₁ is NO₂, R₂ is H, and X-Y is CH₂-S; or

R₁ and R₂ are methoxy, and X-Y is CH₂-S; or

R₁ is H, R₂ is methyl, and X-Y is S(O)-CH₂.

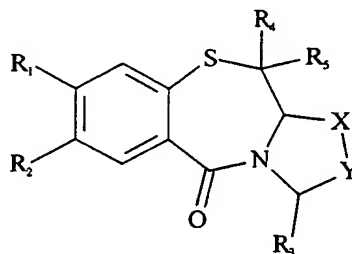
- 45 -

24. The method of claim 21, wherein the compound is



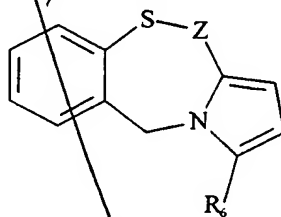
wherein X-Y is S-CH₂ or CH₂-S.

25. The method of claim 21, wherein the compound is



wherein X-Y is CH₂-CH₂;
R₁, R₂, R₃ and R₄ are H; and
R₅ is N(CH₂CH₂)₂NCH₃.

26. The method of claim 21, wherein the compound comprises



wherein R₆ is H and Z is C=O.

27. The method of claim 1, wherein
A is thiazole, benzene, naphthalene, pyridine, pyrimidine, pyrazine, or
quinoline;

R is one or more of halogen or NO₂;

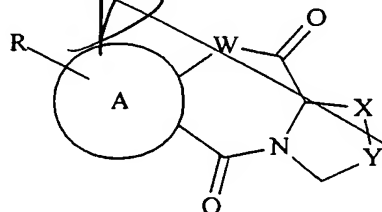
X-Y is CH₂-S, S-CH₂, CH₂-O, CH₂-S(O), S(O)-CH₂, CH₂-CH₂, CH₂-CH₂-
CH₂, or CH₂-CH₂-CH₂-CH₂;

R₄ is H or hydroxy;

R₅ is H, phenyl, or alkylamine;

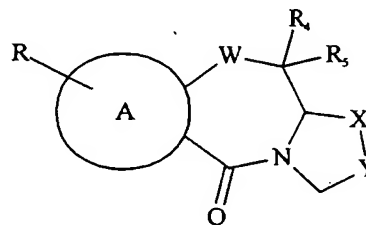
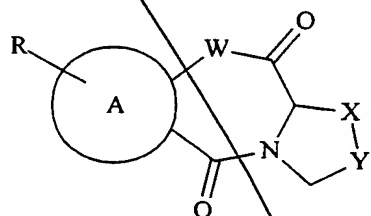
R_6 is H, or substituted or unsubstituted alkyl or amine; and
W is S or O.

28. The method of claim 21, wherein the compound comprises



5 and A is benzene or naphthalene;
R is H, NO_2 , or lower alkoxy; and
X-Y is $\text{CH}_2\text{-S}$ or S-CH_2 .

29. A compound having the following formula, or a pharmaceutically
10 acceptable salt thereof:



wherein A is thiazole, benzene, naphthalene, pyridine, pyrimidine,
pyrazine, or quinoline;

R is one or more of halogen or NO_2 ;

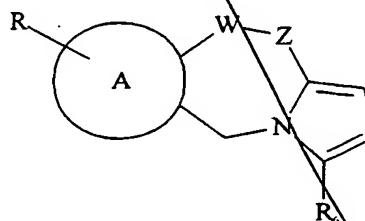
X-Y is $\text{CH}_2\text{-S}$, S-CH_2 , $\text{CH}_2\text{-O}$, $\text{CH}_2\text{-S(O)}$, S(O)-CH_2 , $\text{CH}_2\text{-CH}_2$, $\text{CH}_2\text{-CH}_2\text{-CH}_2$,
15 CH_2 , or $\text{CH}_2\text{-CH}_2\text{-CH}_2\text{-CH}_2$;

R_4 is H or hydroxy;

R_5 is H, phenyl, or alkylamine; and

W is S or O.

20 or wherein the compound is



wherein

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A is thiazole, benzene, naphthalene, pyridine, pyrimidine, pyrazine, or quinoline; and

R is one or more of halogen or NO₂;

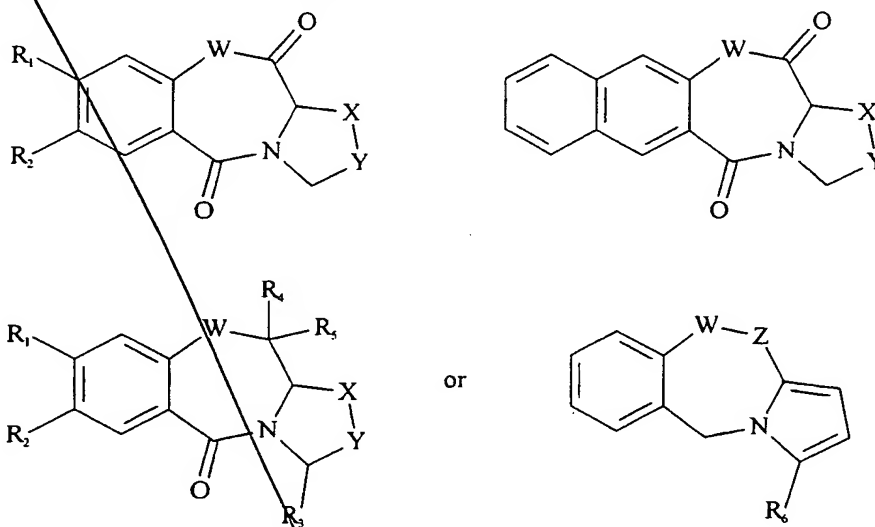
R₆ is H, substituted or unsubstituted alkyl or amine;

W is S or O; and

Z is S, O, CH₂, CH₂CH₂, or C=O.

5

30. A compound having the following formula, or a pharmaceutically acceptable salt thereof, wherein the compound is:



10

wherein

X-Y is S-CH₂, CH₂-S, S(O)-CH₂, CH₂-S(O), or CH₂CH₂;

W is S or O;

R₁ is H or NO₂;

R₂ is H, halogen, lower alkyl or lower alkoxy;

15

R₃ is H;

R₄ is hydroxy or H;

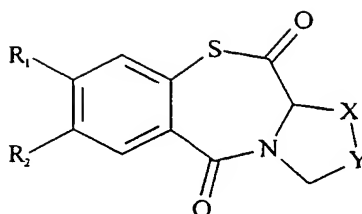
R₅ is phenyl or N(CH₂CH₂)₂NCH₃; and

R₆ is CH₂N(CH₂CH₂)₂NCH₃,

provided that R₁ and R₂ are not both H or not both alkoxy.

20

31. The compound of claim 30, wherein the compound is



and R_1 is H or NO_2 ;

R₂ is H, halogen, lower alkyl or lower alkoxy;

provided that R₁ and R₂ are not both H or not both alkoxy.

5 32. The compound of claim 30, wherein

R_1 is H, R_2 is ~~Cl~~, X-Y is S-CH₂; or

R_1 is H, R_2 is Br, X-Y is S-CH₂; or

R_1 is H, R_2 is CH_3 , X-Y is S- CH_2 ; or

R_1 is H, R_2 is H, $X-Y$ is CH_2-S ; or

10 R_1 is H, R_2 is Cl, $X-Y$ is CH_2-S ; or

R_1 is H, R_2 is Br, X-Y is $\text{CH}_2\text{-S}$; or

R_1 is H, R_2 is CH_3 , $X-Y$ is CH_2-S ; or

R_1 is NO_2 , R_2 is H, $X-Y$ is $\text{CH}_2\text{-S}$; or

R_1 is H, R_2 is OCH_3 , $X-Y$ is CH_2-S ; or

15 R_1 is H, R_2 is H, X-Y is ~~CH₂-O~~; or

R_1 is H, R_2 is CH_3 , X-Y is $S(O)-CH_2$; or

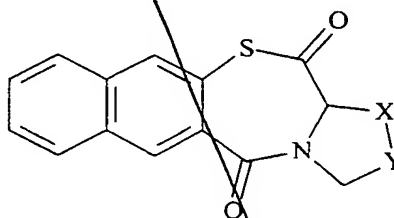
R_1 is H, R_2 is H, X-Y is $\text{CH}_2\text{-S(O)}$; or

R_1 is H, R_2 is Cl, X-Y is $\text{CH}_2\text{-S(O)}$; or

R_1 is H, R_2 is OCH_3 , X-Y is $CH_2-S(O)$.

20

33. The compound of claim 30, wherein the compound is

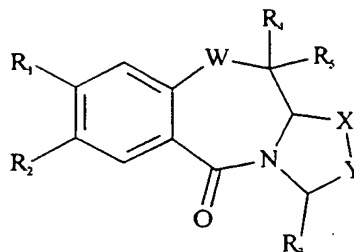
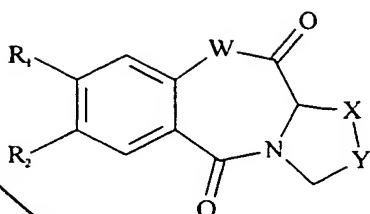


and X-Y is S-CH₂ or CH₂-S.

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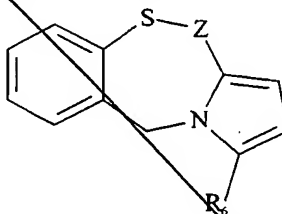
34. The compound of claim 30, wherein X-Y is S-CH₂.

35. The compound of claim 30, wherein the compound is:



and R_1 , R_2 and R_3 are H, R_4 is OH or H;
 R_5 is Ph or $N(CH_2CH_2)_2CH_3$; and
 X-Y is CH_2-CH_2 .

- 5 36. The compound of claim 30, wherein the compound is



and R_6 is $CH_2N(CH_2CH_2)_2NCH_3$.

- 10 37. A pharmaceutical composition comprising the compound of claim
 29, or the pharmaceutically acceptable salt, and a pharmaceutically acceptable
 carrier.

- 15 38. A pharmaceutical composition comprising the compound of claim
 30, or the pharmaceutically acceptable salt, and a pharmaceutically acceptable
 carrier.

- 20 39. A method of screening for an anti-HIV integrase drug, comprising:
 providing an assay of HIV integrase inhibition; and
 using the assay to screen for drugs comprising analogs or derivatives of
 any of the compounds of claim 1.

40. The method of claim 39, wherein the assay detects a thiazepine
 compound that inhibits human immunodeficiency virus type-1 integrase (HIV-1
 IN).

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41. The method of claim 40, further comprising detecting a thiazepine having no detectable effect on reverse transcriptase, protease, and virus attachment.
- 5 42. The method of claim 39, wherein the compound is a thiazolothiazepine.
43. The compound of claim 29, for use in a pharmaceutical composition for the inhibition of HIV integrase.
- 10 44. The compound of claim 43, for use in the treatment of HIV infection.
45. The compound of claim 44, for use as a prophylactic treatment
- 15 against HIV infection.

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